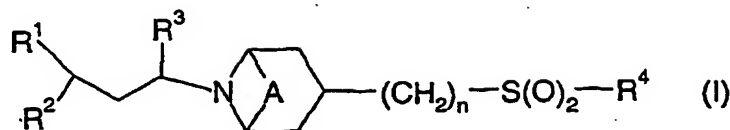


CLAIMS

1. A compound of formula (I):



5 wherein:

A is absent or is (CH₂)₂;

R¹ is C(O)NR¹⁰R¹¹, C(O)₂R¹², NR¹³C(O)R¹⁴, NR¹⁵C(O)NR¹⁶R¹⁷, NR¹⁸C(O)₂R¹⁹, heterocyclyl, aryl or heteroaryl;

R¹⁰, R¹³, R¹⁵, R¹⁶ and R¹⁸ are hydrogen or C₁₋₆ alkyl;

10 R¹¹, R¹², R¹⁴, R¹⁷ and R¹⁹ are C₁₋₈ alkyl (optionally substituted by halo, hydroxy, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, C₃₋₆ cycloalkyl (optionally substituted by halo), C₅₋₆ cycloalkenyl, S(C₁₋₄ alkyl), S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), heteroaryl, aryl, heteroaryloxy or aryloxy), aryl, heteroaryl, C₃₋₇ cycloalkyl (optionally substituted by halo or C₁₋₄ alkyl), C₄₋₇ cycloalkyl fused to a phenyl ring, C₅₋₇ cycloalkenyl, or, heterocyclyl (itself optionally substituted by oxo, C(O)(C₁₋₆ alkyl), S(O)_k(C₁₋₆ alkyl), halo or C₁₋₄ alkyl); or R¹¹, R¹², R¹⁴ and R¹⁷ can also be hydrogen; or R¹⁰ and R¹¹, and/or R¹⁶ and R¹⁷ may join to form a 4-, 5- or 6-membered ring which optionally includes a nitrogen, oxygen or sulphur atom, said ring being optionally substituted by C₁₋₆ alkyl, S(O)_k(C₁₋₆ alkyl) or C(O)(C₁₋₆ alkyl);

20 R² is phenyl, heteroaryl or C₃₋₇ cycloalkyl;

R³ is H or C₁₋₄ alkyl;

R⁴ is heterocyclyl;

n is 1, 2 or 3;

25 aryl, phenyl and heteroaryl moieties are independently optionally substituted by one or more of halo, cyano, nitro, hydroxy, OC(O)NR²⁰R²¹, NR²²R²³, NR²⁴C(O)R²⁵, NR²⁶C(O)NR²⁷R²⁸, S(O)₂NR²⁹R³⁰, NR³¹S(O)₂R³², C(O)NR³³R³⁴, CO₂R³⁶, NR³⁷CO₂R³⁸, S(O)₄R³⁹, OS(O)₂R⁴⁹, C₁₋₆ alkyl (optionally mono-substituted by S(O)₂R⁵⁰ or C(O)NR⁵¹R⁵²), C₂₋₆ alkenyl, C₂₋₆ alkynyl, C₃₋₁₀ cycloalkyl, C₁₋₆ haloalkyl, C₁₋₆ alkoxy(C₁₋₆)alkyl, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, phenyl, phenyl(C₁₋₄)alkyl, phenoxy, phenylthio, phenylS(O), phenylS(O)₂, phenyl(C₁₋₄)alkoxy, heteroaryl,

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heteroaryl(C₁₋₄)alkyl, heteroaryloxy or heteroaryl(C₁₋₄)alkoxy; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, S(C₁₋₄ alkyl), S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), S(O)₂NH₂, S(O)₂NH(C₁₋₄ alkyl), S(O)₂N(C₁₋₄ alkyl)₂, cyano, C₁₋₄ alkyl, C₁₋₄ alkoxy, C(O)NH₂,
 5 C(O)NH(C₁₋₄ alkyl), C(O)N(C₁₋₄ alkyl)₂, CO₂H, CO₂(C₁₋₄ alkyl), NHC(O)(C₁₋₄ alkyl), NHS(O)₂(C₁₋₄ alkyl), CF₃ or OCF₃;

unless otherwise stated heterocyclyl is optionally substituted by C₁₋₆ alkyl [optionally substituted by phenyl {which itself optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, OCF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)} or heteroaryl {which itself optionally substituted
 10 by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}], phenyl {optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, OCF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}, heteroaryl {optionally substituted
 15 by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}, S(O)₂NR⁴⁰R⁴¹, C(O)R⁴², C(O)₂(C₁₋₆ alkyl) (such as tert-butoxycarbonyl), C(O)₂(phenyl(C₁₋₂ alkyl)) (such as benzyloxycarbonyl), C(O)NHR⁴³, S(O)₂R⁴⁴, NHS(O)₂NHR⁴⁵, NHC(O)R⁴⁶,
 20 NHC(O)NHR⁴⁷ or NHS(O)₂R⁴⁸, provided none of these last four substituents is linked to a ring nitrogen;

k, l and q are, independently, 0, 1 or 2;

R²⁰, R²², R²⁴, R²⁶, R²⁷, R²⁹, R³¹, R³³, R³⁷, R⁴⁰ and R⁵¹ are, independently, hydrogen or C₁₋₆ alkyl;

R²¹, R²³, R²⁵, R²⁸, R³⁰, R³², R³⁴, R³⁶, R³⁸, R³⁹, R⁴¹, R⁴², R⁴³, R⁴⁴, R⁴⁵, R⁴⁶, R⁴⁷, R⁴⁸,

25 R⁴⁹, R⁵⁰ and R⁵² are, independently, C₁₋₆ alkyl (optionally substituted by halo, hydroxy, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, C₃₋₆ cycloalkyl, C₅₋₆ cycloalkenyl, S(C₁₋₄ alkyl), S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), heteroaryl, phenyl, heteroaryloxy or phenyloxy), C₃₋₇ cycloalkyl, phenyl or heteroaryl; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, S(C₁₋₄ alkyl),
 30 S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), S(O)₂NH₂, S(O)₂NH(C₁₋₄ alkyl), S(O)₂N(C₁₋₄ alkyl)₂, cyano, C₁₋₄ alkyl, C₁₋₄ alkoxy, C(O)NH₂, C(O)NH(C₁₋₄ alkyl), C(O)N(C₁₋₄ alkyl)₂, CO₂H, CO₂(C₁₋₄ alkyl), NHC(O)(C₁₋₄ alkyl), NHS(O)₂(C₁₋₄ alkyl), C(O)(C₁₋₄ alkyl), CF₃ or OCF₃;

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R^{21} , R^{23} , R^{25} , R^{28} , R^{30} , R^{34} , R^{35} , R^{36} , R^{41} , R^{42} , R^{43} , R^{45} , R^{46} , R^{47} and R^{52} may additionally be hydrogen;

or a pharmaceutically acceptable salt thereof or a solvate thereof.

5 2. A compound as claimed in claim 1 wherein R^1 is $NR^{13}C(O)R^{14}$, wherein R^{13} and R^{14} are as defined in claim 1.

3. A compound as claimed in claim 1 or 2 wherein R^1 is optionally substituted aryl or optionally substituted heteroaryl, wherein the optional substituents are as recited in
10 claim 1.

4 A compound as claimed in claim 1, 2 or 3 wherein R^1 is optionally substituted heterocyclyl.

15 5. A compound as claimed in any one of the preceding claims wherein R^2 is phenyl optionally substituted by halo or CF_3 .

6. A compound as claimed in any one of the preceding claims wherein R^3 is hydrogen.

20 7. A compound as claimed in any one of the preceding claims wherein R^4 is heterocyclyl optionally substituted by oxo, halogen, cyano, hydroxy, C_{1-6} alkyl (itself optionally substituted by halogen, hydroxy, cyano or C_{1-4} alkoxy), C_{2-4} alkenyl, $CO_2(C_{1-4}$ alkyl), $S(O)_2(C_{1-4}$ alkyl), $CH(O)$, $S(O)_2(C_{1-4}$ haloalkyl), $C(O)(C_{1-4}$ alkyl), $C(O)(C_{3-6}$ cycloalkyl), $N(C_{1-4}$ alkyl) $_2$, $C(O)NH_2$, $C(O)N(C_{1-4}$ alkyl) $_2$ or $NHC(O)(C_{1-4}$ alkyl).

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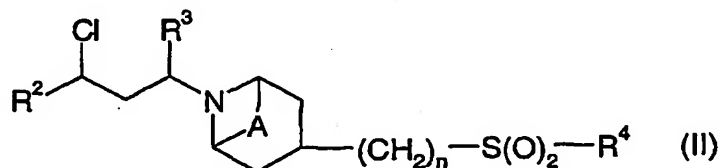
8. A compound as claimed in any one of the preceding claims wherein heterocyclyl is piperidinyl, homopiperazinyl, thiomorpholinyl, pyrrolidinyl, piperazinyl, 1,2,3,6-tetrahydropyridinyl, morpholinyl, 2,5-dihydropyrrolyl, azetidiny, 1,4-oxepanyl, 3-azabicyclo[3.2.1]octan-3-yl, 8-azaspiro[4.5]decanyl or 3-azabicyclo[3.1.0]hex-3-yl.

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9. A compound as claimed in any one of the preceding claims wherein A is absent.

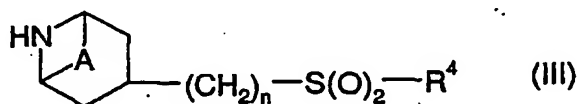
10. A compound as claimed in any one of the preceding claims wherein n is 2.

11. A process for preparing a compound as claimed in claim 1, the process comprising:
- when R^1 is an N-linked optionally substituted heterocycle, reacting a compound of formula (II):

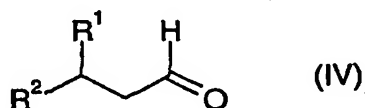


wherein R^2 , R^3 , R^4 , n , A and X are as defined in claim 1, with a compound $R^1\text{H}$ (wherein the H is on a heterocycle ring nitrogen atom) wherein R^1 is as defined in claim 1, in the presence of a suitable base and in a suitable solvent;

- when R^3 is hydrogen, coupling a compound of formula (III):

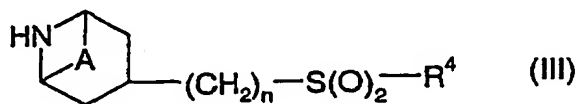


wherein R^4 , n , A and X are as defined in claim 1, with a compound of formula (IV):

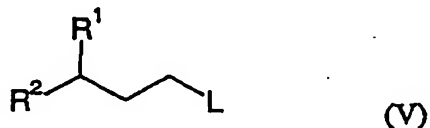


wherein R^1 and R^2 are as defined in claim 1, in the presence of $\text{NaBH}(\text{OAc})_3$ (wherein Ac is $\text{C}(\text{O})\text{CH}_3$) in a suitable solvent at room temperature; or,

- when R^3 is hydrogen, coupling a compound of formula (III):



wherein R^4 , n , A and X are as defined in claim 1, with a compound of formula (V):



wherein R^1 and R^2 are as defined in claim 1 and L is an activated leaving group, in the presence of a base, in a suitable solvent at a temperature from 60°C up to the boiling point of the solvent.

12. A pharmaceutical composition which comprises a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, and a pharmaceutically acceptable adjuvant, diluent or carrier.

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13. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, for use as a medicament.

14. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, in the manufacture of a medicament for use in therapy.

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15. A method of treating a CCR5 mediated disease state comprising administering to a patient in need of such treatment an effective amount of a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof.

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